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**PRV**PATENT- OCH REGISTRERINGSVERKET  
Patentavdelningen

SE 00/1495 10/030924

REC'D 19 SEP 2000

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(71) Sökande CMA/Microdialysis AB, Solna SE  
Applicant (s)

(21) Patentansökningsnummer 9902695-7  
Patent application number

(86) Ingivningsdatum 1999-07-14  
Date of filing

Stockholm, 2000-09-14

För Patent- och registreringsverket  
For the Patent- and Registration Office

Åsa Dahlberg  
Åsa Dahlberg

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Postadress/Adress  
Box 5055  
S-102 42 STOCKHOLM

Telefon/Phone  
+46 8 782 25 00  
Vx 08-782 25 00

Telex  
17978  
PATOREG S

Telefax  
+46 8 666 02 86  
08-666 02 86

## Improved probe

### FIELD OF THE INVENTION

The invention relates to a microdialysis probe. Microdialysis probes of this kind are  
5 described in SE-C-434 214, US,A,5,735,832 and US,A,5,741,284.

The meaning of specific wordings in this text should be interpreted as follows:

The word probe should be interpreted also as catheter.

The inlet and outlet of the probe as described may in case of a reversed flow be used  
10 as outlet and inlet, respectively.

Perfusion liquid is the liquid used in the microdialysis, which is allowed to enter the  
probe and there take up substances from the surrounding tissue through a membrane.

The perfusion liquid becomes the dialysate after the dialysis.

Non-invasive investigating examination in this document refers to e.g. X-ray, NMR  
15 or the like techniques.

### BACKGROUND OF THE INVENTION

Microdialysis is a method of examination in which a probe is inserted into tissue in  
vivo, such that one side of a semi-permeable membrane is in contact with tissue and  
20 extra cellular liquid and the other side is flushed or rinsed with a dialysis liquid  
(perfusate) which takes-up substances from the extra cellular liquid through the  
membrane. These substances can then be analyzed in the dialysate on or after exiting  
the probe.

25 Microdialysis probes are by nature fragile and very small, which requires great care  
in inserting and withdrawing the probe from the tissue in which it is used. However,  
it is also of great importance that the probe when inserted into tissue of a living  
person, is placed in the intended location such that when measuring the probe  
actually through microdialysis measures the intended chemical/biological variables  
30 that at each measurement is of interest. It is of course important in these  
measurements to know exactly what is measured.

The use of microdialysis becoming more frequent and common raises other problems such as monitoring and control of the probe during insertion and use. It is a fact that microdialysis provides a unique possibility to examine the equilibria of substances and/or the amounts present or missing of substances or to monitor specific changes in the status of substances connected with e.g. the use of medicaments, in surgery etc.

The monitoring and control of the probe position during insertion/withdrawal and use has been an obstacle in so far that the smallness and the material of the probe does not make possible the use of common methods for detecting the probe once the insertion has been started. This becomes more problematic the deeper into the tissue the microdialysis is to take place.

#### 15 SUMMARY OF THE INVENTION.

It is thus an object of the invention is to provide a microdialysis probe, the location of which may be monitored and controlled using means such as X-rays or the like during insertion/withdrawal or during dialysis in order to facilitate the placement of the probe at a predetermined location and to control the location of the probe.

20

It is also an object of the invention to provide a microdialysis probe, which is suitable for the general use in living tissue when taking samples for e.g. diagnostic purposes.

25 In accordance with the invention, these and other objects evident from the description of the invention are accomplished in a microdialysis probe in that characterized by presence of a position indicating object imparting such characteristics to the distal part of the probe as to allow non-invasive examination of the location of the distal part of the microdialysis probe.

**BRIEF DESCRIPTION OF THE DRAWINGS.**

The invention will now be described by way of example and with reference to the accompanying drawings in which:

5 Fig. 1 shows, partially in section, a first embodiment of a microdialysis probe in section according to the invention.

Fig. 2 shows a detailed view of the foremost part of the probe according to the embodiment as shown in Fig. 1.

10

Fig. 3 a - c shows in detail the sealing of the most distal part of the first tube according to the embodiment shown in Fig. 1 and Fig. 2.

15 Fig. 4 shows a second embodiment of a microdialysis probe in section according to the invention.

**DETAILED DESCRIPTION OF PREFERRED FORMS OF THE INVENTION.**

A first embodiment of the microdialysis probe according to the invention is shown in Fig. 1 and Fig 2. The probe exhibits a distal end piece 110 comprised of glue which holds and seals a plug 130 within the distal part of a membrane 115. This comprises the foremost tip of the probe. The membrane 115 is preferably tubular. A proximal tubular fitting 111 and a proximal end piece 113 comprises the other end of the probe as such. The proximal tubular fitting 111 is permanently fastened to a proximal end piece 113. The proximal end of the membrane 115 is fastened to the proximal tubular fitting 111.

20 In the proximal end piece 113 two tubes 107 and 108 constituting the inlet to the probe and the outlet from the probe are connected to the probe, such as to let the perfusion liquid pass through the same. Note in the definitions above the possibility of reversed flow.

30

Within the membrane 115, which is in the form of a tube made from semi-permeable material, and also within the proximal tubular fitting 111, a first tube 116 extends essentially from the proximal end of the probe to the distal end. The first tube 116 is closed at the most distal end by a position indicating object, a plug 130, and exhibits at least one aperture 117 at or near the distal end. The aperture 117 constitutes a passage for the perfusion liquid entering the space 118 defined by the first tube 116 and the dialysis membrane 115 in combination with the proximal tubular fitting 111 and the distal tubular fitting 112. For the withdrawal of the perfusion liquid a second tube 119 extends from the proximal end of the probe and opens up into the same space 118 somewhere near to the proximal end of the probe thereby forming an exit for the perfusion liquid. The perfusion liquid has now become a dialysate having acquired substances exchanged over the semi-permeable membrane. The distal end piece 110 of the probe may e.g. be fastened in a permanent way to the distal end of the first tube 116.

15

To give a proper understanding of the invention, exemplary dimensions are given here. The length of the probe may be e.g. 5 cm from the most distal end of the same to the proximal part of the proximal tubular fitting 111. The length of the tubular fitting may be approximately 2 cm, thus the length of the membrane may be approximately 3 cm. The diameter of the proximal tubular fitting may be approximately 1 mm and the outer diameter of the membrane may be approximately 0.6 mm.

20

The plug 130 shown in Fig. 2 may preferably be manufactured from gold. Other materials may of course be used but the reason for using gold is the ductility and the relative opaqueness for X-rays and also the chemical and physiological inertness exhibited by gold.

25

In a second embodiment the plug 130 may be made such as to make the distal end of the microdialysis probe visible during examination using NMR (Nuclear Magnetic Resonance).

30

This plug could e.g. have the form of a hollow amorphous plug filled with air or the like, which would impart characteristics to the plug such that it will be possible to locate the plug using NMR.

5

One way of accomplishing the sealing at the distal end of the first tube 316 is shown in Fig 3 a - c.

10 In Fig. 3a) the form of an exemplary first tube 316 according to the invention is shown. The tube has been subjected to corbelling in order to widen the diameter of the same enough to accommodate the sealing plug 330.

In Fig. 3 b) a preferred embodiment of the sealing plug 330 is shown and in Fig. 3c) the first tube 316 with the sealing plug 330 in place is shown.

15

A second embodiment of the microdialysis probe according to the invention is shown in Fig. 4. The probe exhibits a distal end piece 410 and a distal tubular fitting 412. The distal tubular fitting 412 in combination with the end piece 410 comprises the foremost tip of the probe. A proximal tubular fitting 411 and a proximal end piece 413 comprises the other end of the probe as such. The proximal tubular fitting 411 is permanently fastened to a proximal end piece 413. A membrane 415 is fastened to the distal tubular fitting 412, the membrane 415 having a smaller diameter than the fitting. The membrane is preferably tubular. The fitting itself being closed at the most distal end thereof e.g. by using glue or the like, forming the distal end 410. The other end of the membrane 415 is fastened to the proximal tubular fitting 411.

25

In order to accomplish an X-ray opaqueness of the foremost part of the probe the distal end may be formed as a half-sphere or the like from a material which is opaque to X-rays or the X-ray opaqueness creating material may be incorporated in the glue as such.

30



In the end proximal piece 413 two tubes 407 and 408 constituting the inlet to the probe and the outlet from the probe are connected to the probe, such as to let the perfusion liquid pass through the same. Note above the possibility of reversed flow.

5

In this embodiment the dimensions of the probe will be approximately the same as in the embodiment described in connection with Fig 1.

Within the membrane 415, which is in the form of a tube made from semi-permeable material, first tube 416 extends essentially from the proximal end of the probe to the distal end. The first tube 416 has a closed distal end and has at least one aperture 417 at or near the distal end. The aperture 417 constitutes a passage for the perfusion liquid entering the space 418 defined by the first tube 416 and the dialysis membrane 415 in combination with the proximal tubular fitting 411 and the distal tubular fitting 412. For the withdrawal of the perfusion liquid a second tube 419 extends from the proximal end of the probe and opens up into the same space 418 somewhere near to the proximal end of the probe thereby forming an exit for the perfusion liquid.

20 In the same manner as has been described in connection with the first embodiment of the invention the end piece 410 sealing distal end of the probe may be made such as to make the same visible during examination using NMR (Nuclear Magnetic Resonance). This could e.g. be done by incorporating in the sealing glue a hollow amorphous plug filled with air or the like, which would impart characteristics to the distal end of the probe such that it will be possible to locate the probe end by using NMR.

The invention has been described under reference to embodiments of the same. It should be understood that the above describes an exemplary embodiment of the distal end of the probe itself and the constructive details thereof may vary within the scope of the claims or be independent of the constructive details of the distal end of

the probe depending on different embodiments of the invention. The scope of the invention however is described by the appended claims.

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Claims

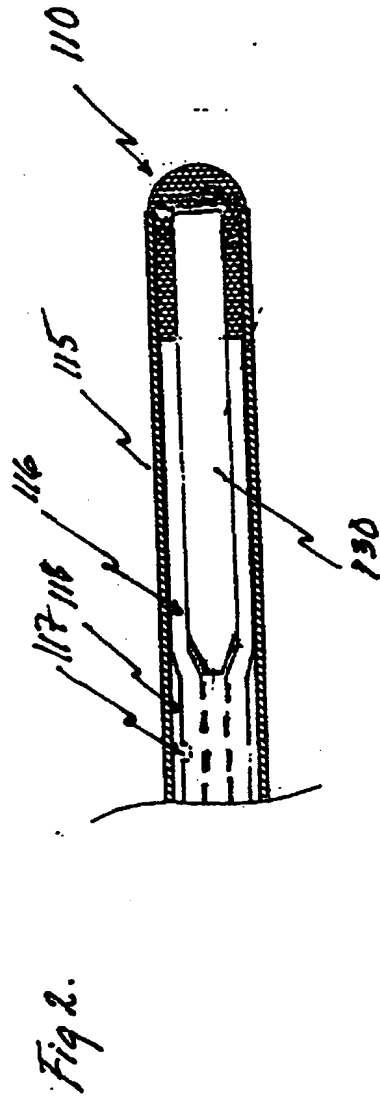
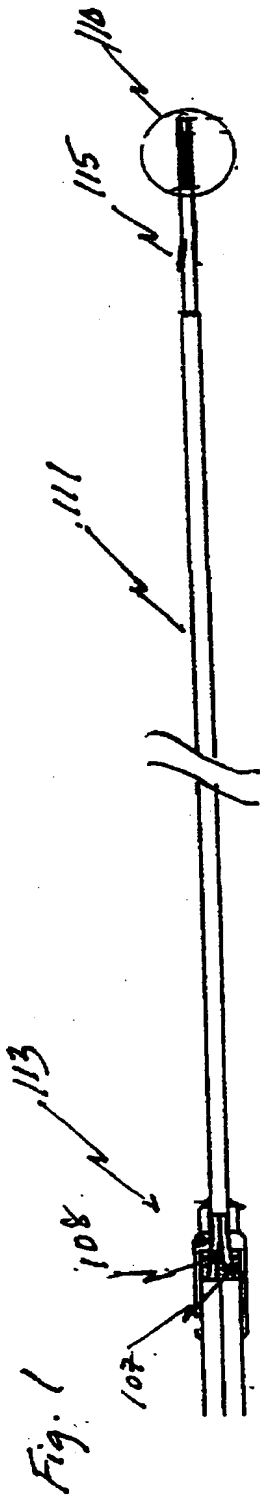
1. A microdialysis probe, comprising a dialysis membrane (115,215) located and supported between a closed distal end of the probe and a proximal end of the same, said membrane (115,215) essentially surrounding a space (118,218) for passage of perfusion liquid; said probe having inlet and outlet means (107,108; 207,208) for perfusion liquid, characterized by presence of a position indicating object (130,410) imparting such characteristics to the distal part of the probe as to allow non-invasive examination of the location of the distal part of the microdialysis probe.
2. A microdialysis probe according to claim 1, characterized in said position indicating object being an x-ray opaqueness giving object.
3. A microdialysis probe according to claim 1, characterized in said position indicating object being an object visible when using NMR (Nuclear Magnetic Resonance).
4. A microdialysis probe according to any preceding claim, characterized in a first tube for inlet of perfusion liquid into the space between the membrane and the first tube, said first tube exhibiting a through-hole for the passage of the perfusion liquid near the distal end of the first tube, said most distal end of the first tube being sealed by a plug of said position indicating object.
5. A microdialysis probe according to claim 1, 2 or 4, characterized in that said position indicating object is a plug made of gold.
6. A microdialysis probe according to claim 1, 2 or 4, characterized in that said position indicating object is a plug made of any bio-compatible metal, alloy or amorphous compound which imparts the x-ray opaqueness necessary.

7. A microdialysis probe according to claim 1, 2 or 4, characterized in that said position indicating object is a plug made of compound dispersed in the glue sealing the probe which imparts the x-ray opaqueness to the plug.
- 5 8. A microdialysis probe according to claim 1 or 3, characterized in that said position indicating object is a hollow object filled with air or the like.

Abstract

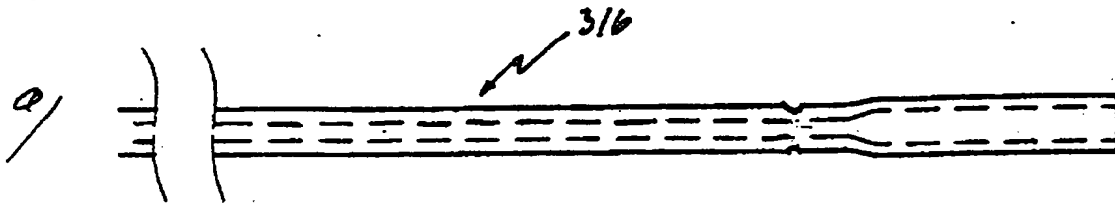
The invention concerns a microdialysis probe, comprising a dialysis membrane (115,415) located and supported between a closed distal end of the probe and a proximal end of the same, said membrane (115,415) essentially surrounding a space (118,418) for passage of perfusion liquid; said probe having inlet and outlet means (107,108; 407,408) for perfusion liquid, characterized by presence of a position indicating object imparting such characteristics to the distal part of the probe as to allow non-invasive examination of the location of the distal part of the microdialysis probe.

Fig. 2

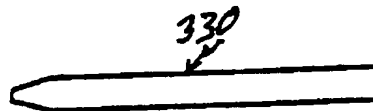


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Fig. 3



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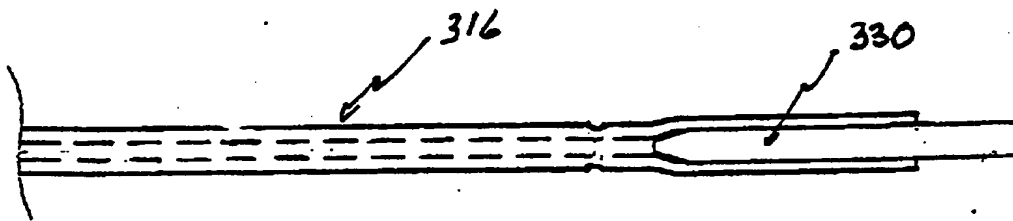


Fig. 4

